

JORDAN AND HAMBURG LLP  
122 East 42nd Street,  
New York, NY 10168  
Tel: (212) 986-2340 Fax: (212) 953-7733  
Customer No.: 000028107

Docket No.: F-7333  
Filing Date: February 22, 2002

**Certificate of Express Mailing Under 37 CFR 1.10**

"Express Mail" label number EV 021 743 560 US Date of Deposit 2/22/2002

I hereby certify that this correspondence is being deposited with the United States Postal Service "Express Mail Post Office to Addressee" service under 37 CFR 1.10 in an envelope addressed to ASSISTANT COMMISSIONER OF PATENTS, WASHINGTON, DC 20231 having the above "Express Mail" label number on the date of deposit indicated above

Madeline Gonzalez  
(Typed or printed name of person mailing paper or fee)

*Madeline Gonzalez*  
(Signature of person mailing paper or fee)

THE ASSISTANT COMMISSIONER FOR PATENTS  
Washington, D. C. 20231

- ☐ ATTN: BOX PATENT APPLICATION  
☐ ATTN: BOX DESIGN PATENT APPLICATION  
☒ ATTN: BOX PCT  
☐ ATTN: BOX PROVISIONAL PATENT APPLICATION  
☒ THIS IS THE 35 U.S.C 371 NATIONAL STAGE OF PCT/JP00/06040 FILED

September 6, 2000

Sir:

Transmitted herewith for filing is the ☒ Utility ☐ Design ☒ nonprovisional ☐ provisional  
patent application of:

Inventor / Application Identifier: Setsuko UEDA, et al.

☐ See Inventor Information Sheet attached

For: **CHEMICALLY PEELING AGENT**

- ☐ This is a new patent application.  
☒ This is the 35 U.S.C. 371 National Stage Application of the above-identified PCT Application.  
☐ This is a provisional patent application.  
☐ This is a: ☐ Continuation Application  
☐ Divisional Application  
☐ Continuation-in-Part Application  
of prior Application Serial No. \_\_\_\_  
☐ Cancel in this application original claims \_\_\_\_ of the prior application before  
calculating the filing fee.  
☐ Amend the specification by inserting before the first line the sentence:  
-- This is a ☐ Continuation, ☐ Divisional, ☐ Continuation-in-part, of  
Application

☐ Incorporation By Reference. The entire disclosure of the prior application, from which a copy of the oath or declaration is supplied, is considered as being part of the disclosure of the accompanying application and is hereby incorporated by reference therein.

ENCLOSED ARE THE FOLLOWING:		
	0	Sheets of drawings ( <input type="checkbox"/> formal <input type="checkbox"/> informal size A4 ).
X	11	Pages of specification including abstract and claims.
X	11	Total pages.
X	Combined Declaration and Power of Attorney	
	X	Newly executed
		Copy from prior application
		Inventors deleted; see attached statement
	Sequence Listing	
		Computer Readable Copy
		Paper copy
		Statement verifying identity of above copies
X	Return Receipt Postcard	
X	Preliminary Amendment	
	Assignment to:	
		Assignment is of record in prior application Serial No. .
		Assignment Recordation Form Cover Sheet.
		Charge \$40.00 to Deposit Account No. 10-1250 for recording Assignment.
X	Information Disclosure Statement	
X	Information Disclosure Citation	
	English translation	
X	Application Data Sheet	

PRIORITY CLAIMS	
	Applicant hereby claims the benefit of the filing date of the following provisional application(s) under the provision of 35 USC 119.
X	<p>Applicant hereby claims the benefit under the provisions of 35 USC 119 of the filing dates of the following applications as indicated below:</p> <p style="margin-left: 40px;">Japan Patent Appln. No. 11-251802, filed September 6, 1999, Priority Claimed</p> <p style="margin-left: 40px;">Japan Patent Appln. No. 2000-155339, filed May 25, 2000, Priority Claimed</p> <p style="margin-left: 40px;">of which certified copies thereof</p>
	will follow
	are enclosed
X	have been filed in the International Bureau
	were filed in prior application:

CLAIMS FILED AND FILING FEE CALCULATION					
ITEM	---			Rate	Applied Fee
<input type="checkbox"/> Base Fee - Non PCT	---			\$740	
<input type="checkbox"/> Base Fee - PCT IPEA-US	---			\$710	
<input type="checkbox"/> Base Fee - PCT ISA-US	---			\$740	
<input type="checkbox"/> Base Fee - PCT not ISA or IPEA	---			\$1,040	
<input checked="" type="checkbox"/> Base Fee - PCT with EPO or JPO Search Report	---			\$890	\$890
<input type="checkbox"/> Base Fee - Design	---			\$330	
<input type="checkbox"/> Base Fee - Provisional	---			\$160	
Claim Fees	Number Filed	Base Number	Number Extra over Base	---	
Total Claims	15	20	0	\$18	\$0
Independent Claims	1	3	0	\$84	\$0
Multiple Dependent Claim Fee	---			\$280	\$280
<input checked="" type="checkbox"/> Small Entity Status is Asserted	---				(\$585)
<input type="checkbox"/> Foreign Language Filing Fee	---			\$130	\$0
<b>TOTAL FILING FEE</b>					<b>\$585</b>

- [X] Please charge Deposit Account No. 10-1250 in the amount of the above TOTAL FILING FEE. A duplicate copy of this sheet is attached.
- [X] Please charge to Deposit Account No. 10-1250 any further fees due for filing or during prosecution of this application under: 37 CFR 1.16; 37 CFR 1.17; and 37 CFR 1.492.

JORDAN AND HAMBURG LLP

By 

C. Bruce Hamburg  
Reg. No. 22,389  
Attorney for Applicant

F-7333

**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

Applicant : Setsuko UEDA, et al.  
Serial No. : U.S. National Stage of PCT/JP00/06040  
Filed September 6, 2000  
Filed : Concurrently herewith  
For : CHEMICALLY PEELING AGENT  
Group Art Unit : (Not yet known)  
Examiner : (Not yet known)

Hon. Commissioner of Patents  
and Trademarks  
Washington, D.C. 20231

**PRELIMINARY AMENDMENT**

Sir:

Preliminary to examination, please amend this application as follows:

**IN THE SPECIFICATION:**

Amend the paragraph on page 2, lines 10-22, as follows, the changes  
being shown by underlining in an Appendix hereto:

F-7333

In order to achieve the object as described above, the present invention provides a chemically peeling agent comprising a component (A) having the following general formula:



(wherein B is an alcohol residue;

AO is an alkylene-oxy group having 3 to 18 carbon atoms;

a is an integer of 1 or more;

m is an integer of 4 or more; and

n is 0 or an integer of 1 or more;

provided that a molar amount m of the oxidized ethylene to be added is a value that amounts to 40% or more of the entire molecular weight of an ethylene oxide chain moiety)

and a phenol compound (B) formulated in an amount ranging from 10% to 30% by weight.

(Please note that in line 10 of the above amended paragraph there is an underlined character which is part of the original specification.)

F-7333

Amend the third complete paragraph at page 4 of the specification as follows, the changes being shown by brackets and underlining in an Appendix hereto:

The amount of the phenol compound to be added may be in the range of from 10 to 30% by weight. When the phenol compounds are to be combined with two kinds or more, the amounts of the compounds are to be appropriately selected within the scope so as to effectively peel the horny layer off and cause no side effects.

Delete the heading and four paragraphs at page 7, lines 6-21 of the specification.

Amend the heading at page 7, line 22 of the specification as follows, the changes being shown by brackets and underlining in the Appendix hereto:

#### Example 4

Delete the heading and six paragraphs at page 8, lines 4-23 of the specification.

F-7333

Amend the heading at page 8, line 24 of the specification, the changes being shown by brackets and underscoring in an Appendix hereto:

Example 5

Delete the heading and five paragraphs at page 9, lines 6-21, of the specification.

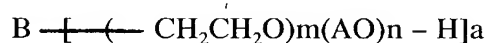


F-7333

**IN THE CLAIMS:**

Amend claims 1 and 5-7 as follows, the changes being shown by brackets and underlining in an Appendix hereto:

1. (Amended) A chemically peeling agent comprising a component (A) having the following general formula:



(wherein B is an alcohol residue;

AO is an alkylene-oxy group having 3 to 18 carbon atoms;

a is an integer of 1 or more;

m is an integer of 4 or more; and

n is 0 or an integer of 1 or more;

provided that a molar amount m of the oxidized ethylene to be added is a value that amounts to 40% or more of the entire molecular weight of an ethylene oxide chain moiety)

and a phenol compound (B) formulated in an amount ranging from 10% to 30% by weight.

5. (Amended) The chemically peeling agent as claimed in claim 1, 2 or 3, wherein said phenol compound (B) is salicylic acid.

F-7333

6. (Amended) The chemically peeling agent as claimed in claim 1, 2 or 3, wherein said phenol compound (A) is a member selected from the group consisting of a phenol and a resorcinol.

7. (Amended) The chemically peeling agent as claimed in claim 1, 2 or 3, wherein a gelling agent or a viscosity-adjusting agent is added to the component (A) and the phenol compound (B).

**IN THE ABSTRACT:**

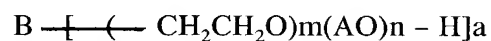
Substitute for the original Abstract the Abstract appended hereto on a separate page.

The amendments to the original Abstract are shown by underlining in an Appendix hereto.

F-7333

## ABSTRACT

The chemically peeling agent contains a component (A) having the following general formula:



(wherein B is an alcohol residue;

AO is an alkylene-oxy group having 3 to 18 carbon atoms;

a is an integer of 1 or more;

m is an integer of 4 or more; and

n is 0 or an integer of 1 or more;

provided that a molar amount  $\underline{m}$  of the oxidized ethylene to be added is a value that amounts to 40% or more of the entire molecular weight of an ethylene oxide chain moiety)

and a phenol compound (B) formulated in an amount ranging from 10% to 30% by weight.

F-7333

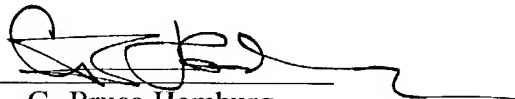
**REMARKS**

This Amendment essentially corresponds to an Article 34 Amendment filed in the International Stage. In addition, claim dependencies are amended so that no multiple dependent claim depends from another multiple dependent claim.

Respectfully submitted,

Jordan and Hamburg LLP

By



C. Bruce Hamburg

Reg. No. 22,389

Attorney for Applicants

122 East 42nd Street  
New York, New York 10168  
(212) 986-2340

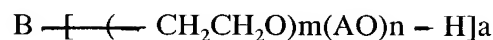
CBH/pb

F-7333

## APPENDIX I

AMENDED CLAIMS WITH AMENDMENTS INDICATED THEREIN  
BY BRACKETS AND UNDERLINING

1. (Amended) A chemically peeling agent comprising a component (A) having the following general formula:



(wherein B is an alcohol residue;

AO is an alkylene-oxy group having 3 to 18 carbon atoms;

a is an integer of 1 or more;

m is an integer of 4 or more; and

n is 0 or an integer of 1 or more;

provided that a molar amount m of the oxidized ethylene to be

added is a value that amounts to 40% or more of the entire

molecular weight of an ethylene oxide chain moiety)

and a phenol compound (B) formulated in an amount ranging from 10% to 30% by weight.

5. (Amended) The chemically peeling agent as claimed in [any one of claims 1, 2, 3 and 4] claim 1, 2 or 3, wherein said phenol compound (B) is salicylic acid.

F-7333

6. (Amended) The chemically peeling agent as claimed in [any one of claims 1, 2, 3 and 4] claim 1, 2 or 3, wherein said phenol compound (A) is a member selected from the group consisting of a phenol and a resorcinol.

7. (Amended) The chemically peeling agent as claimed in [any one of claims 1, 2, 3, 4, 5 and 6] claim 1, 2 or 3, wherein a gelling agent or a viscosity-adjusting agent is added to the component (A) and the phenol compound (B).

F-7333

## APPENDIX II

AMENDED SPECIFICATION PARAGRAPHS WITH AMENDMENTS  
INDICATED THEREIN BY BRACKETS AND UNDERLINING

Page 2, the paragraph consisting of lines 10-22, is amended as indicated below:

In order to achieve the object as described above, the present invention provides a chemically peeling agent comprising a component (A) having the following general formula:



(wherein B is an alcohol residue;

AO is an alkylene-oxy group having 3 to 18 carbon atoms;

a is an integer of 1 or more;

m is an integer of 4 or more; and

n is 0 or an integer of 1 or more;

provided that a molar amount m of the oxidized ethylene to be added is a value that amounts to 40% or more of the entire molecular weight of an ethylene oxide chain moiety)

and a phenol compound (B) formulated in an amount ranging from 10% to 30% by weight.

F-7333

Page 4, 3rd full paragraph, is amended as indicated below:

The amount of the phenol compound to be added may be in the range of from [0.01 % to 70% by weight in the case of salicylic acid, from 0.01 % to 60% by weight in the case of phenol, and from 0.01 % to 50% by weight in the case of resorcinol] 10 to 30% by weight. When the phenol compounds are to be combined with two kinds or more, the amounts of the compounds are to be appropriately selected within the scope so as to effectively peel the horny layer off and cause no side effects.

Page 7: after the 1st full paragraph, delete the heading and delete the 2nd, 3rd, 4th, and 5th full paragraphs, as indicated below:

[Example 4:

A chemically peeling agent was prepared by mixing 60% by weight of polyethylene glycol 1500 with 40% by weight of salicylic acid.



F-7333

A face of each of 10 women in the sixties was pasted with the chemically peeling agent and the agent was wiped out in 3 minutes. The pasting was carried out once a month for three months.

An observation was conducted by means of a digital camera and the observation revealed a disappearance of shallow wrinkles and a rise in hue and brightness for all the persons under test. An observation by a scanning electronic microscope of cheek replica indicates an apparent disappearance of wrinkles in the skin. A comparison before and after treatment with the chemically peeling agent indicated a light increase of the water content in the horny layer, a light rise in the amount of vaporization of moisture through the skin, and improvements in development of the skin (when observed with a cuticle meter).

Further, no side effects including, for example, red spots, pigmentation, scars and so on were recognized for all the persons under test.]

Page 7:           after 5th full paragraph, amend the heading as indicated  
below:

Example [5] 4:

F-7333

Page 8: delete the heading before the 1st full paragraph and delete the 1st, 2nd, 3rd, 4th, 5th and 6th full paragraphs, as indicated below:

[Example 6:

A chemically peeling agent was prepared by mixing 50% by weight of polyethylene glycol 1500 with 50% by weight of phenol.

A face of each of 10 women in the sixties was pasted with the chemically peeling agent and the agent was wiped out in 3 minutes. The pasting was carried out once a month for three months.

An observation was conducted by means of a digital camera and the observation revealed a disappearance of shallow wrinkles and a rise in hue and brightness for all the persons under test. An observation by a scanning electronic microscope of cheek replica indicates an apparent disappearance of wrinkles in the skin.

A comparison before and after the treatment with the chemically peeling agent indicated a light increase of the water content in the horny layer, a light rise in the amount of vaporization of moisture through the skin, and improvements in development of the skin (when observed with a cuticle meter).

F-7333

Although the chemically peeling agent contained phenol in such a concentration as high as 50% by weight, no side effects including, for example, red spots, pigmentation, scars and so on were recognized for all the persons under test.

It was also found that the chemically peeling agent containing phenol in the concentration of 60% by weight in polyethylene glycol exhibited the effects similar to the agent containing 50% by weight of phenol.]

Page 8: the heading after the 6th paragraph, is amended as indicated below:

Example [7] 5:

Page 9: Delete the heading before the 2nd full paragraph and delete the 2nd, 3rd, 4th, 5th and 6th paragraphs, as indicated below:

[Example 8:

A chemically peeling agent was prepared by mixing 50% by weight of polyethylene glycol 1500 with 50% by weight of resorcinol.

F-7333

A face of each of 10 women in the sixties was pasted with the chemically peeling agent and the agent was wiped out in 10 minutes. The pasting was carried out once a month for three months.

An observation was conducted by means of a digital camera and the observation revealed a disappearance of shallow wrinkles and a rise in hue and brightness for all the persons under test. An observation by a scanning electronic microscope of cheek replica indicates an apparent disappearance of wrinkles in the skin.

A comparison before and after the treatment with the chemically peeling agent indicated a light increase of the water content in the horny layer, a light rise in the amount of vaporization of moisture through the skin, and improvements in development of the skin (when observed with a cuticle meter).

Further, no side effects including, for example, red spots, pigmentation, scars and so on were recognized for all the persons under test.]

F-7333

## APPENDIX III

AMENDED ABSTRACT WITH AMENDMENTS INDICATED  
THEREIN BY BRACKETS AND UNDERLINING

## ABSTRACT

[The present invention provides a chemically peeling agent that can exhibit the efficacy of the main component thereof without causing less side effects.]

The chemically peeling agent contains a component (A) having the following general formula:



(wherein B is an alcohol residue;

AO is an alkylene-oxy group having 3 to 18 carbon atoms;

a is an integer of 1 or more;

m is an integer of 4 or more; and

n is 0 or an integer of 1 or more;

provided that a molar amount m of the oxidized ethylene to be added is a value that amounts to 40% or more of the entire molecular weight of an ethylene oxide chain moiety)

F-7333

and a phenol compound (B) formulated in an amount ranging from 10% to 30%  
by weight.

## DESCRIPTION

## CHEMICALLY PEELING AGENT

## TECHNICAL FIELD

The present invention relates to a chemically peeling agent adapted to remove wrinkles, spots (pigmentation such as geriatric pigment spots) and a somber color on the skin, to treat pimples, and to improve a greasy skin. The term "chemically peeling agent" referred to in this description is intended to mean a chemically peeling agent for use as medicine or cosmetics.

In the West, removal of wrinkles, spots and so on is considered as one of methods for medical treatments. A treatment method that is generally adopted at hospitals of dermatology, orthopedics, or cosmetic surgery in the western countries, involves pasting the skin with an aqueous solution of different concentrations of a chemically peeling agent, including trichloroacetic acid (TCA), phenol and so on, for an appropriate duration of time to artificially make a chemical burn (corrosion) and thereafter to reproduce normal cells on the skin in a natural way.

This treatment with such chemically peeling agents, however, is effective for white people, it may cause the skin of Asian people problems with side effects including red spots, pigmentation, scars and so on after operation.

Recently, it has been found that  $\alpha$ -hydroxy acid (AHA) is relatively safe and it is effective for the peeling treatment for Asian people. This treatment has now been conducted as a general treatment method. The effects and the side effects to be produced by this method, however, greatly depend upon the concentration and pH of AHA, so that it suffers from the difficulty that this treatment method requires experienced skills.

More recently, the treatment with a solution of salicylic acid in an alcohol has been adopted in the U. S. and it has been found to be effective for white people. This treatment method, however, causes Asian people problems with severe side effects including, for example, flare or pain during treatment and pigmentation after treatment.

As a result of extensive and long-lasting research, it was found by the present inventor that, although an agent containing salicylic acid in polyethylene glycol cannot be absorbed through the skin so that it has been considered to be ineffective for the treatment of skin diseases, a mixture of salicylic acid with polyethylene glycol allows salicylic acid to be sustained in the polyethylene glycol and retained in the horny layer without penetrating into a sebaceous matter at a high concentration and to strongly peel off the horny layer only without causing any systemic side effects. Further, it was found that a mixture of a phenol compound, such as phenol or resorcinol, with a

polyethylene glycol or an equivalent compound could exhibit the effects similar to the mixture of salicylic acid with the polyethylene glycol.

The present invention has been completed on the basis of these findings and has the object to provide a chemically peeling agent that does not cause any side effects including red spots, pigmentation and scars after operation, refreshes the skin (removing fine wrinkles and providing the skin with flexibility), removes spots, and improve a somber color on the skin. Further, the chemically peeling agent can treat pimples and improve a greasy skin.

#### DISCLOSURE OF INVENTION

In order to achieve the object as described above, the present invention provides a chemically peeling agent comprising a component (A) having the following general formula:



(wherein B is an alcohol residue;

AO is an alkylene-oxy group having 3 to 18 carbon atoms;

a is an integer of 1 or more;

m is an integer of 4 or more; and

n is 0 or an integer of 1 or more;

provided that a molar amount  $\underline{m}$  of the oxidized ethylene to be added is a value that amounts to 40% or more of the entire molecular weight of an ethylene oxide chain moiety)

and a phenol compound (B).

The component (A) to be used for the present invention may be represented by the above general formula.

In the above general formula, the alcohol to be represented by reference symbol B is intended to mean a mono-valent alcohol including, for example, an alkyl alcohol such as methanol, ethanol, butanol, lauryl alcohol, myristyl alcohol, palmityl alcohol, stearyl alcohol, cetyl alcohol, etc., and an alkenyl alcohol such as linoleyl alcohol, palmitoyl alcohol, oleyl alcohol, etc., a di-valent alcohol such as ethylene glycol, propylene glycol, etc., a tri-valent alcohol such as glycerin, trimethylol propane, triethanol amine, etc., a tetra-valent alcohol such as pentaerythritol, diglycerin, etc. There may also be used other poly-valent alcohols such as sorbitol, polyglycerin and so on.

The alkylene-oxy group having 3 to 18 carbon atoms, as referred to by reference symbol AO, may include, for example, propylene-oxy, butylene-oxy,



tetrahydrofuran,  $\alpha$ -olefin-oxy, and so on. The alkylene-oxy groups having 3 and 4 carbon atoms, such as oxido-propylene, oxido-butylene and tetrahydrofuran, are preferred.

In the above general formula, reference symbol "a" is an integer of 1 or more. When the alcohol to be used for the present invention is a mono-valent alcohol, the reference symbol "a" is 1. When the alcohol to be used therefor is a di-valent alcohol, the reference symbol "a" is 2. Likewise, when the alcohol to be used therefor is a tri-valent alcohol, the reference symbol "a" is 3. Further, when the alcohol to be used therefor is a poly-valent alcohol, the reference symbol "a" is the integer corresponding to the valence of the alcohol used.

In the above general formula, reference symbol "m" is intended to mean an average molar amount of ethylene oxide to be added. The number of a polymerization chain of the ethylene oxide has to be at least 4.

Reference symbol "n" is intended to mean an average molar amount of an oxidized alkylene to be added. The number of a polymerization chain of the oxidized alkylene is zero or 1 or more.

The manner of polymerization of the ethylene oxide and the alkylene oxide is random or block polymerization.

The molar amount  $\underline{m}$  of the ethylene oxide to be added is set to amount to 40% or more of the entire molecular weight of the ethylene oxide chain. This setting is based on the fact that, if the molar amount  $\underline{m}$  of the ethylene oxide to be added would be less than the above molar amount, the phenol compound such as salicylic acid would become unlikely to be sustained in the polyethylene glycol derivative.

The component (A) may be synthesized in a conventional manner, for example, by reacting the ethylene oxide and the alkylene oxide with the alkyl alcohol or the alkenyl alcohol under an inert gas such as nitrogen or the like in the presence of a basic catalyst such as sodium hydroxide, potassium hydroxide or the like or an acidic catalyst such as boron tetrafluoride, tin tetrachloride or the like.

Specific examples of the compounds (A) may include, for example, a polyethylene glycol and a derivative thereof, a polyoxyethylene polyoxypropylene glycol and a derivative thereof, and a polyoxyethylene long-chain alkyl ether and a polyoxyethylene long-chain alkenyl ether to be used as a non-ionic surfactant. Among these examples, the polyethylene glycol is listed as trade name "Macrogol" in the Japanese Pharmacopoeia and the Regulations of Medicinal Additives. Likewise, a block copolymer of ethylene oxide or propylene oxide is listed as a poloximer therein. The phenol compound to be added as the main component can be sustained in those compounds and retained in the horny layer without penetrating into a sebaceous matter at a high concentration. Therefore, these compounds can preferably be used because

they have no risk of causing any systemic side effects and can peel only the horny layer off. These compounds can be used singly or in combination of two or more.

As the phenol compound (B) to be used for the present invention, there may be mentioned phenol, resorcinol, salicylic acid and so on.

Among the phenol compounds (B), salicylic acid is preferred. The phenol compounds may be used singly or in combination of two or more.

The amount of the phenol compound to be added may be in the range of from 0.01% to 70% by weight in the case of salicylic acid, from 0.01% to 60% by weight in the case of phenol, and from 0.01% to 50% by weight in the case of resorcinol. When the phenol compounds are to be combined with two kinds or more, the amounts of the compounds are to be appropriately selected within the scope so as to effectively peel the horny layer off and cause no side effects.

The chemically peeling agent according to the present invention may preferably include an alkyl acrylate-methacrylate copolymer in the amount of from 0.1% to 5% by weight as a gelling agent or a viscosity-adjusting agent in order to prevent the softening of the product particularly in the summer season.

To the chemically peeling agent according to the present invention, there may be added various conventional additives for use with an ointment or cosmetics, which do not interfere with the efficacy or impose any influence upon the efficacy of the chemically peeling agent. Such additives may include, for example, an aromatic, a surfactant, a preservative, an anti-oxidant, a moisturizing agent, and so on, and they are to be used in an appropriate amount that does not reduce the efficacy of the chemically peeling agent. In addition, vitamin A acid may be added. Moreover, the addition of the surfactant and so on can preferably improve the efficacy of the agent.

The chemically peeling agent according to the present invention may be prepared in a conventional way, for example, by mixing the component (A) in a molten state with the component (B) at ambient temperature or elevated temperature or under addition of pressure and adding various additives thereto as needed.

The chemically peeling agent according to the present invention may be applied, for example, by pasting the chemically peeling agent on the skin and wiping the agent out from the skin after a given period of time. The application of this agent can serve as removing the epidermis (mainly the cuticle) of the skin and imposing influences upon the cells of the stratum spinosum epidermidis and the stratum basale epidermidis of the epidermis, thereby causing the reproduction of the fibroblast of the corium. The aged corium portion can be replaced with the reproduced fibroblast to induce the skin-restoring effects. This can remove wrinkles on the skin and restore the elastic power in the skin. At the same time, the cuticle of the hair follicle is also peeled off and the accumulated cuticle can be removed, thereby curing pimples. The disinfecting effects of

the main components of the chemically peeling agent can synergistically act on the pimples and improve the greasy skin, too.

As the chemically peeling agent according to the present invention can re-structure the corium of the skin by peeling the horny layer off and reproducing the fibroblast in the corium, melanin withering the curium can be removed from the focus as time elapses. Further, the fibrous tissues reproduced in layers on top of melanocyte of a neoblast so that bluish to brownish color in hue can also be masked when looked at an appearance. Therefore, chromatosis induced by those causes can also be cured so that spots and dark or somber color on the skin can be caused to disappear.

The duration for pasting the chemically peeling agent may be preferably set to be for from 3 minutes to 20 minutes, although it is not restricted to the particular period of time. It can be appropriately selected from the duration of time that does not cause any side effects and can effectively produce the peeling effects.

It is to be noted herein that, if the phenol compound such as salicylic acid is applied at a low concentration, the chemically peeling agent can be preferably applied to the skin after removal of the cuticle, whereby the cuticle remaining in the hair follicle or in the skin can be removed effectively without causing any side effects.

#### BEST MODES FOR CARRYING OUT THE INVENTION

The present invention will be described in more detail by way of examples. In the following description, examples are described in which polyethylene glycol is used as the component (A). It is to be noted herein, however, that the use of the other components (A) can also exhibit the effects substantially identical to or similar to the examples where polyethylene glycol is used as the component (A). Further, it is to be noted herein that, although polyethylene glycol having a molecular weight of 1,500 is used in each of the following examples, a polyethylene glycol having a molecular weight ranging from 1,500 to 20,000 can also be used preferably from the point of view of penetration of the main component into the skin tissue and the unlikelihood of an occurrence of side effects or from other reasons. Moreover, such polyethylene glycols may also be used singly or in combination of two or more while adjusting viscosity, penetrating ability and other properties.

##### Example 1:

A chemically peeling agent was prepared by mixing 90% by weight of polyethylene glycol 1500 with 10% by weight of salicylic acid.

A face of each of 20 women in the sixties was pasted with the chemically peeling agent and the agent was wiped out in 20 minutes. The pasting was carried out once a month for three months. It should be noted herein that salicylic acid used herein is the one that is generally used as a softening agent for the horny layer of the skin or a

disinfectant.

An observation was conducted by means of a digital camera. The observation revealed a slight disappearance of shallow wrinkles and a slight rise in hue and brightness for all the persons under test. An observation by a scanning electronic microscope of cheek replica indicates a slight disappearance of wrinkles in the skin.

Further, no side effects including, for example, red spots, pigmentation, scars and so on were recognized for all the persons under test.

It was also found that the agent containing salicylic acid in the concentration of 10% by weight or less could remove the horny layer left in the hair follicle and the skin effectively without causing any side effects, when the skin was pasted with the agent after removal of the horny layer.

Example 2:

A chemically peeling agent was prepared by mixing 80% by weight of polyethylene glycol 1500 with 20% by weight of salicylic acid.

A face of each of 50 women in the sixties was pasted with the chemically peeling agent and the agent was wiped out in 10 minutes. The pasting was carried out once a month for three months.

An observation was conducted by means of a digital camera and the observation revealed a disappearance of shallow wrinkles and a rise in hue and brightness for all the persons under test. An observation by a scanning electronic microscope of cheek replica indicates an apparent disappearance of wrinkles in the skin. A comparison before and after the treatment with the chemically peeling agent indicated an increase of the water content in the horny layer, a rise in the amount of vaporization of moisture through the skin, and improvements in development of the skin (when observed with a cuticle meter).

Further, no side effects including, for example, red spots, pigmentation, scars and so on were recognized for all the persons under test.

Example 3:

A chemically peeling agent was prepared by mixing 70% by weight of polyethylene glycol 1500 with 30% by weight of salicylic acid.

A face of each of 10 women in the sixties was pasted with the chemically peeling agent and the agent was wiped out in 5 minutes. The pasting was carried out once a month for three months.

An observation was conducted by means of a digital camera and the observation revealed a disappearance of shallow wrinkles and a rise in hue and brightness for all the persons under test. An observation by a scanning electronic microscope of cheek replica indicates an apparent disappearance of wrinkles in the skin. A comparison before and after the treatment with the chemically peeling agent

indicated a light increase of the water content in the horny layer, a light rise in the amount of vaporization of moisture through the skin, and improvements in development of the skin (when observed with a cuticle meter).

Further, no side effects including, for example, red spots, pigmentation, scars and so on were recognized for all the persons under test.

Example 4:

A chemically peeling agent was prepared by mixing 60% by weight of polyethylene glycol 1500 with 40% by weight of salicylic acid.

A face of each of 10 women in the sixties was pasted with the chemically peeling agent and the agent was wiped out in 3 minutes. The pasting was carried out once a month for three months.

An observation was conducted by means of a digital camera and the observation revealed a disappearance of shallow wrinkles and a rise in hue and brightness for all the persons under test. An observation by a scanning electronic microscope of cheek replica indicates an apparent disappearance of wrinkles in the skin. A comparison before and after the treatment with the chemically peeling agent indicated a light increase of the water content in the horny layer, a light rise in the amount of vaporization of moisture through the skin, and improvements in development of the skin (when observed with a cuticle meter).

Further, no side effects including, for example, red spots, pigmentation, scars and so on were recognized for all the persons under test.

Example 5:

A chemically peeling agent was prepared by mixing 90% by weight of polyethylene glycol 1500 with 10% by weight of phenol.

A face of each of 10 women in the sixties was pasted with the chemically peeling agent and the agent was wiped out in 3 minutes. The pasting was carried out once a month for three months.

An observation was conducted by means of a digital camera and the observation revealed a disappearance of shallow wrinkles and a rise in hue and brightness for all the persons under test. An observation by a scanning electronic microscope of cheek replica indicates an apparent disappearance of wrinkles in the skin. A comparison before and after the treatment with the chemically peeling agent indicated a light increase of the water content in the horny layer, a light rise in the amount of vaporization of moisture through the skin, and improvements in development of the skin (when observed with a cuticle meter).

Further, no side effects including, for example, red spots, pigmentation, scars and so on were recognized for all the persons under test.

It was also found that the agent containing phenol in the concentration of 5%

by weight or less could remove the horny layer left in the hair follicle and the skin effectively without causing any side effects, when the skin was pasted with the agent after removal of the horny layer.

Example 6:

A chemically peeling agent was prepared by mixing 50% by weight of polyethylene glycol 1500 with 50% by weight of phenol.

A face of each of 10 women in the sixties was pasted with the chemically peeling agent and the agent was wiped out in 3 minutes. The pasting was carried out once a month for three months.

An observation was conducted by means of a digital camera and the observation revealed a disappearance of shallow wrinkles and a rise in hue and brightness for all the persons under test. An observation by a scanning electronic microscope of cheek replica indicates an apparent disappearance of wrinkles in the skin.

A comparison before and after the treatment with the chemically peeling agent indicated a light increase of the water content in the horny layer, a light rise in the amount of vaporization of moisture through the skin, and improvements in development of the skin (when observed with a cuticle meter).

Although the chemically peeling agent contained phenol in such a concentration as high as 50% by weight, no side effects including, for example, red spots, pigmentation, scars and so on were recognized for all the persons under test.

It was also found that the chemically peeling agent containing phenol in the concentration of 60% by weight in polyethylene glycol exhibited the effects similar to the agent containing 50% by weight of phenol.

Example 7:

A chemically peeling agent was prepared by mixing 90% by weight of polyethylene glycol 1500 with 10% by weight of resorcinol.

A face of each of 10 women in the sixties was pasted with the chemically peeling agent and the agent was wiped out in 10 minutes. The pasting was carried out once a month for three months.

An observation was conducted by means of a digital camera and the observation revealed a disappearance of shallow wrinkles and a rise in hue and brightness for all the persons under test. An observation by a scanning electronic microscope of cheek replica indicates an apparent disappearance of wrinkles in the skin.

A comparison before and after the treatment with the chemically peeling agent indicated a light increase of the water content in the horny layer, a light rise in the amount of vaporization of moisture through the skin, and improvements in development of the skin (when observed with a cuticle meter).

Further, no side effects including, for example, red spots, pigmentation, scars

and so on were recognized for all the persons under test.

It was also found that the chemically peeling agent containing resorcinol in the concentration of less than 10% by weight in polyethylene glycol could remove the horny layer left in the hair follicle and the skin effectively without causing any side effects, when the skin was pasted with the agent after removal of the horny layer.

Example 8:

A chemically peeling agent was prepared by mixing 50% by weight of polyethylene glycol 1500 with 50% by weight of resorcinol.

A face of each of 10 women in the sixties was pasted with the chemically peeling agent and the agent was wiped out in 10 minutes. The pasting was carried out once a month for three months.

An observation was conducted by means of a digital camera and the observation revealed a disappearance of shallow wrinkles and a rise in hue and brightness for all the persons under test. An observation by a scanning electronic microscope of cheek replica indicates an apparent disappearance of wrinkles in the skin.

A comparison before and after the treatment with the chemically peeling agent indicated a light increase of the water content in the horny layer, a light rise in the amount of vaporization of moisture through the skin, and improvements in development of the skin (when observed with a cuticle meter).

Further, no side effects including, for example, red spots, pigmentation, scars and so on were recognized for all the persons under test.

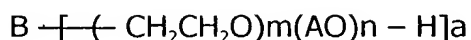
#### INDUSTRIAL UTILIZABILITY

The chemically peeling agent according to the present invention can retain the phenol compound in the component (A) and sustain the phenol compound in the horny layer without penetrating in the sebaceous matter at a high concentration so that it can strongly peel only the horny layer off without causing any risk of an occurrence of systemic side effects. Therefore, the chemically peeling agent according to the present invention can effectively remove wrinkles, spots (pigmentation such as geriatric pigment spots) and a somber color on the skin. It can also be used for treatment of pimples and improvements in the greasy skin.

For the present invention, the polyethylene glycol is used as a substrate so that it can serve as adhering to the skin well upon pasting the skin with the agent without causing any irritating. Further, the agent can be easily dissolved in water so that it can be readily washed away with water. Therefore, the chemically peeling agent according to the present invention can be used effectively and safely without requiring experienced skills.

## CLAIMS

1. A chemically peeling agent comprising a component (A) having the following general formula:



(wherein B is an alcohol residue;

AO is an alkylene-oxy group having 3 to 18 carbon atoms;

a is an integer of 1 or more;

m is an integer of 4 or more; and

n is 0 or an integer of 1 or more;

provided that a molar amount m of ethylene oxide to be added is a value that amounts to 40% or more of the entire molecular weight of an ethylene oxide chain moiety)

and a phenol compound (B).

2. The chemically peeling agent as claimed in claim 1, wherein said component (A) is a polyethylene glycol.

3. The chemically peeling agent as claimed in claim 1, wherein said component (A) is a block polymer of ethylene oxide or propylene oxide.

4. The chemically peeling agent as claimed in claim 1, 2 or 3, wherein the alcohol residue of the component (A) is derived from an alkyl alcohol or an alkenyl alcohol.

5. The chemically peeling agent as claimed in any one of claims 1, 2, 3 and 4, wherein said phenol compound (B) is salicylic acid.

6. The chemically peeling agent as claimed in any one of claims 1, 2, 3 and 4, wherein said phenol compound (A) is a member selected from the group consisting of a phenol and a resorcinol.

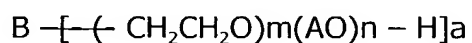
7. The chemically peeling agent as claimed in any one of claims 1, 2, 3, 4, 5 and 6, wherein a gelling agent or a viscosity-adjusting agent is added to the component (A) and the phenol compound (B).



## ABSTRACT

The present invention provides a chemically peeling agent that can exhibit the efficacy of the main component thereof without causing less side effects.

The chemically peeling agent contains a component (A) having the following general formula:



(wherein B is an alcohol residue;

AO is an alkylene-oxy group having 3 to 18 carbon atoms;

a is an integer of 1 or more;

m is an integer of 4 or more; and

n is 0 or an integer of 1 or more;

provided that a molar amount  $\underline{m}$  of the ethylene oxide to be added is a value that amounts to 40% or more of the entire molecular weight of an ethylene oxide chain moiety)

and a phenol compound (B).

# **COMBINED DECLARATION FOR PATENT APPLICATION AND POWER OF ATTORNEY**

(Includes Reference to PCT International Applications)

Attorney's Docket Number

As a below named inventor, I hereby declare that:

My residence, post office address and citizenship are as stated below next to my name,

I believe I am the original, first and sole inventor (if only one name is listed below) or an original, first and joint inventor (if plural names are listed below) of the subject matter which is claimed and for which a patent is sought on the invention entitled:

CHEMICALLY PEELING AGENT

the specification of which (check only one item below):

- ☐ is attached hereto.
- ☐ was filed as United States application  
Serial No. \_\_\_\_\_  
on \_\_\_\_\_,  
and was amended  
on \_\_\_\_\_ (if applicable).
- ☒ was filed as PCT international application  
Number PCT/JP00/06040  
on September 6, 2000  
and was amended under PCT Article 19  
on \_\_\_\_\_ (if applicable).

I hereby state that I have reviewed and understand the contents of the above-identified specification, including the claims, as amended by any amendment referred to above.

I acknowledge the duty to disclose information which is material to the patentability of this application in accordance with Title 37, Code of Federal Regulations, §1.56(a).

I hereby claim foreign priority benefits under Title 35, United States Code, §119 of any foreign application(s) for patent or inventor's certificate or of any PCT international application(s) designating at least one country other than the United States of America listed below and have also identified below any foreign application(s) for patent or inventor's certificate or any PCT international application(s) designating at least one country other than the United States of America filed by me on the same subject matter having a filing date before that of the application(s) of which priority is claimed:

PRIOR FOREIGN/PCT APPLICATION(S) AND ANY PRIORITY CLAIMS UNDER 35 U.S.C. 119:			
Country (if PCT indicate "PCT")	Application Number	Date of Filing (day, month, year)	Priority Claimed Under 35 USC 119
Japan	11-251802	6 September 1999	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No
Japan	2000-155339	25 May 2000	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No
			<input type="checkbox"/> Yes <input type="checkbox"/> No
			<input type="checkbox"/> Yes <input type="checkbox"/> No
			<input type="checkbox"/> Yes <input type="checkbox"/> No

**COMBINED DECLARATION FOR PATENT APPLICATION AND  
POWER OF ATTORNEY (Continued)**  
(Includes Reference to PCT International Applications)

Attorney's Docket Number

I hereby claim the benefit under Title 35, United States Code, §120 of any United States application(s) or PCT international application(s) designating the United States of America that is/are listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in that/those prior application(s) in the manner provided by the first paragraph of Title 35, United States Code, §112, I acknowledge the duty to disclose material information as defined in Title 37, Code of Federal Regulations, §1.56(a) which occurred between the filing date of the prior application(s) and the national or PCT international filing date of this application:

PRIOR U.S. APPLICATIONS OR PCT INTERNATIONAL APPLICATIONS DESIGNATING THE U.S. FOR BENEFIT UNDER 35 U.S.C. 120:					
U.S. APPLICATIONS			STATUS (Check One)		
U.S. Application Number	U. S. Filing Date		Patented	Pending	Abandoned
PCT APPLICATIONS DESIGNATING THE U.S.					
PCT Application No.	PCT Filing Date	U.S. Serial Numbers Assigned (if any)			

POWER OF ATTORNEY: As a named inventor, I hereby appoint the following attorney(s) and/or agent(s) to prosecute this application and transact all business in the Patent and Trademark Office connected therewith.

Frank J. Jordan Reg. No. 20,456  
C. Bruce Hamburg Reg. No. 22,389

Herbert F. Ruschmann Reg. No. 35,341  
Jacqueline M. Steady Reg. No. 44,354  
Derek S. Jessen Reg. No. 48,213  
Lawrence Wechsler Reg. No. 36,049  
Marvin Turken Reg. No. 18,330  
Alfred D'Andrea Reg. No. 27,752

Send Correspondence To: Jordan and Hamburg LLP  
122 East 42nd Street  
New York, New York 10168

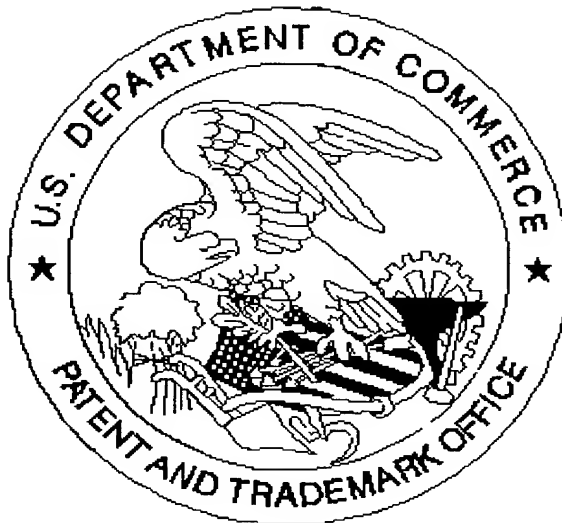
Direct Telephone Calls to:  
C. Bruce Hamburg  
(212) 986-2340

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under section 1001 of Title 18 of the United States Code, and that such willful false statements may jeopardize the validity of the application or any patent issuing thereon.

Full Name of Sole or First Inventor <u>Setsuko UEDA</u>	Inventor's Signature <i>Setsuko Ueda</i>	Date February 5, 2002
Residence <u>Fukuoka-shi, Fukuoka, Japan JPX</u>	Citizenship Japan	
Post Office Address <u>2-3, Kasumigaoka 3-chome, Higashi-ku, Fukuoka-shi, Fukuoka, Japan</u>		

Full Name of Second Joint Inventor, if any <u>Kaori UEDA</u>	Inventor's Signature <i>Kaori Ueda</i>	Date February 5, 2002
Residence <u>Fukuoka-shi, Fukuoka, Japan JPX</u>	Citizenship Japan	
Post Office Address <u>2-3, Kasumigaoka 3-chome, Higashi-ku, Fukuoka-shi, Fukuoka, Japan</u>		

United States Patent & Trademark Office  
Office of Initial Patent Examination -- Scanning Division



Application deficiencies found during scanning:

☐ Page(s) only 2 of recalculation 1 of 3 were not present  
for scanning. (Document title)

☐ Page(s) \_\_\_\_\_ of \_\_\_\_\_ were not present  
for scanning. (Document title)

☐ *Scanned copy is best available.*